

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of:

Schiemann et al.

Group Art Unit: 1625

Serial No.: 10/583,689

Examiner: Rita J. Desai

Filed: Dec. 14, 2004

For: 2-(Hetero)-aryl substituted tetrahydroquinoline

DECLARATION UNDER 37 C.F.R. § 1.132

Honorable Commissioner of
Patents and Trademarks
Washington, D.C. 20231

SIR:

Frank Zenke, being duly warned, deposes and says:

I am a citizen of Germany residing at Darmstadt, Germany;

I am a biologist by training and experience;

the degree of Dr. rer. nat. was bestowed on me by the Heinrich-Heine University of Düsseldorf, Germany in 1995;

from 1995 to 1996 I was employed at the Institute for Microbiology, Heinrich-Heine University of Düsseldorf, Germany and the Institute for Genetics, Martin-Luther University Halle;

from 1997 to 2001 I was postdoctoral scientist at the Department of Immunology, The Scripps Research Institute, La Jolla, USA;

since October 2001 I am an employee at Merck KGaA, Darmstadt, Germany;

I am author or co-author of several papers and patents in the fields of biochemistry and cell biology.

The test results presented below for the present application support that the claimed compounds are inhibitors of the mitotic motor protein Eg5.

I have carried out, or supervised experiments for testing these inhibitors according to the methods described within the genus claimed in the pending application.

Pharmacological Report

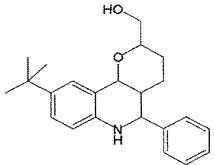
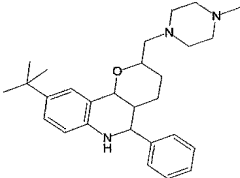
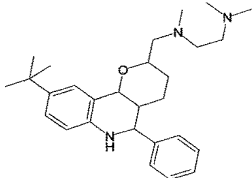
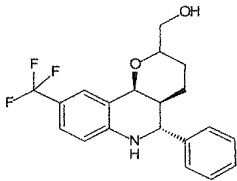
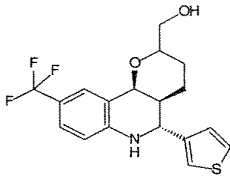
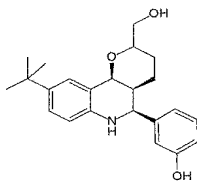
HCT116 cell viability / proliferation assay

The proliferation data have been obtained by means of a conventional assay, which was known to the skilled artisan for the measurement of such biological data at the date of filing this application. Human colon carcinoma cells (HCT116, ATCC CCL 247) were cultured at 37°C and 10% CO₂ in MEM alpha (Invitrogen, #22571) supplemented with 10% (v/v) heat-inactivated fetal bovine serum (Pan Biotech, #3302), 2mM glutamine (Invitrogen, #25030) and 1mM sodium pyruvate (Invitrogen, #11360). Cells were detached from F75 cell culture flasks (Nunc 734 2066) with trypsin/EDTA (Invitrogen, #15400), counted using a Thoma chamber (VWR, #631-1117), and plated in 96-well microtiter plates (Falcon 353227) in fresh medium (cell density of HCT116: 1×10^4 cells in 0.1ml medium per cavity). Cells were cultured overnight and treated on the next day with Eg5 inhibitors or with vehicle alone (0.5% DMSO; Merck, #1.02931) for 48hrs. Wells containing culture medium alone served as background controls.

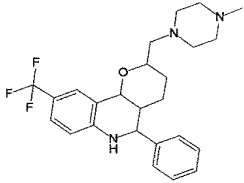
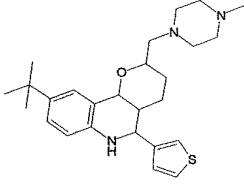
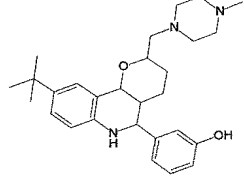
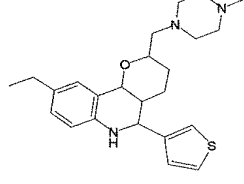
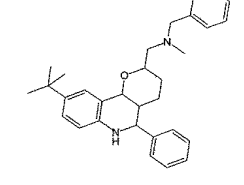
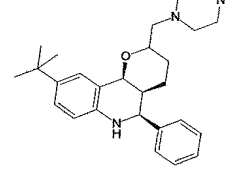
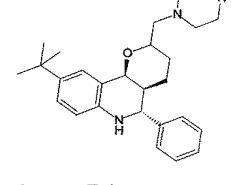
At the end of compound incubation the medium was discarded, the remaining attached cells were washed once with 1x PBS (Life Technologies 14190-094), and stained for 15 minutes with crystal violet solution (0.5% crystal violet, 20% methanol). Subsequently, the wells were washed thoroughly with deionised water until no colour clouds came off the cells. The plates were dried at room temperature overnight or in a 37°C incubator for at least one hour. The bound crystal violet was extracted with 200µl methanol per well and absorption was measured at 550nm wavelength in a spectrophotometer (Tecan Reader SLT Rainbow, Berthold Reader Mithras, LB 940). The mean absorption value of the medium controls was subtracted from all other wells. DMSO control treated cells served as positive control (100%). IC₅₀ values were determined using a curve fitting algorithm of the statistics program RS/1 (Brooks Automation Inc., Massachusetts).

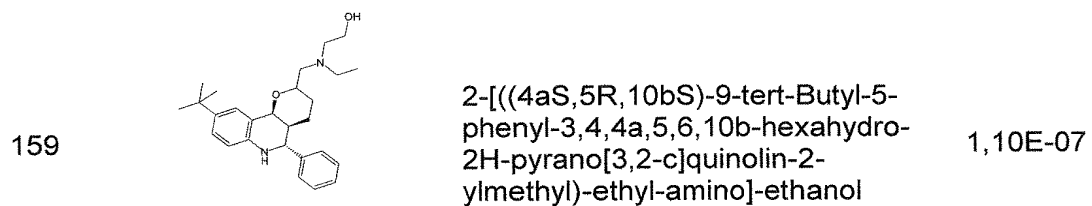
The test data of the Pharmacological Report (Table I) show the compounds tested and their IC₅₀-values. The "no." refers to present application where compounds are disclosed.

Table 1: IC₅₀ values for inhibiting Eg5

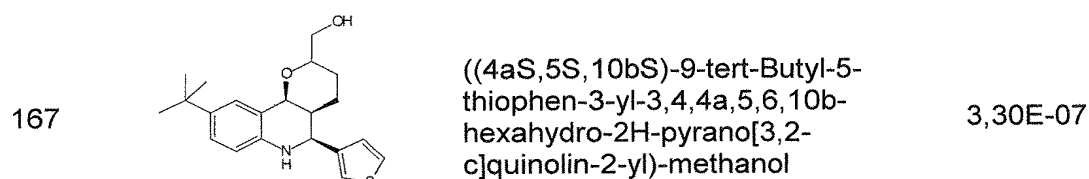
No.	CHEMISTRY	NAME	Proliferation IC ₅₀ [M]
60		(9-tert-Butyl-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	4,30E-08
	Chemistry 37		
89		9-tert-Butyl-2-(4-methyl-piperazin-1-ylmethyl)-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	6,60E-07
	Chemistry 354		
96		N-(9-tert-Butyl-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-N,N',N'-trimethyl-ethane-1,2-diamine	3,90E-07
	Chemistry 346		
120		((4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	1,20E-07
	Chemistry 89		
121		((4aS,5R,10bS)-5-Thiophen-3-yl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	3,42E-07
	Chemistry 88		
122		3-((4aS,5S,10bS)-9-tert-Butyl-2-hydroxymethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-5-yl)-phenol	1,10E-06
	Chemistry 87		

126		((4aS,5R,10bS)-9-tert-Butyl-5-thiophen-3-yl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	2,64E-08
Chemistry 83			
127		3-((4aS,5R,10bS)-2-Hydroxymethyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-5-yl)-phenol	5,80E-08
Chemistry 82			
133		(2S,4aS,5S,10bS)-9-tert-Butyl-2-(4-methyl-piperazin-1-ylmethyl)-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	4,10E-07
Chemistry 76			
137		((2R,4aS,5R,10bS)-9-tert-Butyl-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	2,60E-08
Chemistry 72			
138		(9-tert-Butyl-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	3,04E-08
Chemistry 71			
139		(2S,4aS,5S,10bS)-9-tert-Butyl-2-(4-methyl-piperazin-1-ylmethyl)-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	8,00E-07
Chemistry 70			
145		9-Ethyl-2-(4-methyl-piperazin-1-ylmethyl)-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	1,10E-06
Chemistry 63			

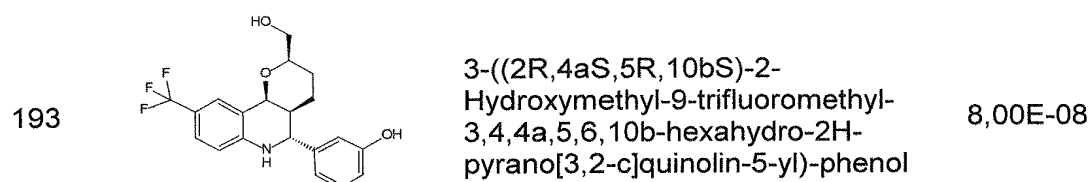
146		2-(4-Methyl-piperazin-1-ylmethyl)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	3,30E-06
	Chemistry 62		
147		9-tert-Butyl-2-(4-methyl-piperazin-1-ylmethyl)-5-thiophen-3-yl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	1,10E-06
	Chemistry 61		
148		3-[9-tert-Butyl-2-(4-methyl-piperazin-1-ylmethyl)-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-5-yl]-phenol	2,40E-07
	Chemistry 60		
150		9-Ethyl-2-(4-methyl-piperazin-1-ylmethyl)-5-thiophen-3-yl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	4,30E-06
	Chemistry 58		
151		Benzyl-(9-tert-butyl-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-methyl-amine	6,70E-06
	Chemistry 57		
153		(4aS,5S,10bS)-9-tert-Butyl-2-(4-methyl-piperazin-1-ylmethyl)-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	2,80E-06
	Chemistry 55		
154		(4aS,5R,10bS)-9-tert-Butyl-2-(4-methyl-piperazin-1-ylmethyl)-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	1,50E-07
	Chemistry 54		



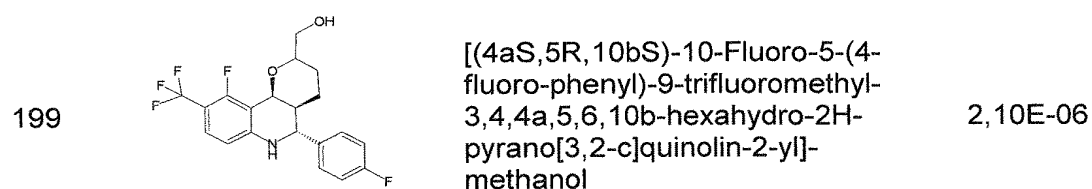
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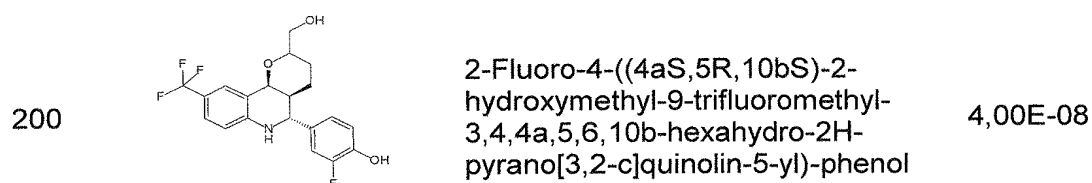
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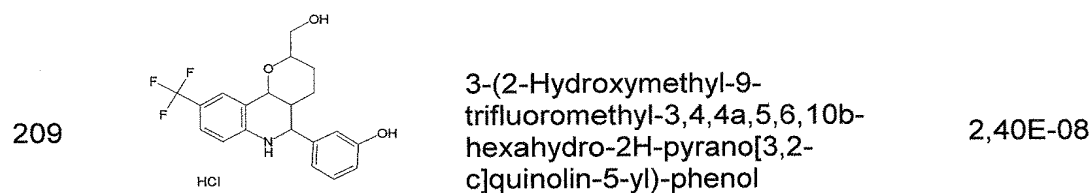
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Chemistry 126

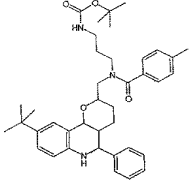
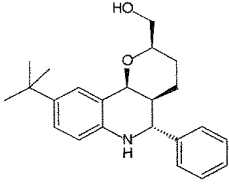
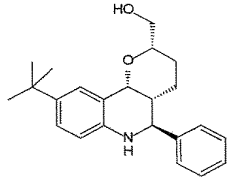
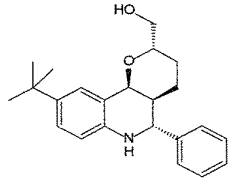
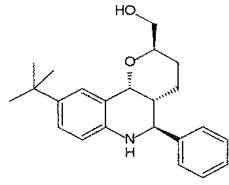
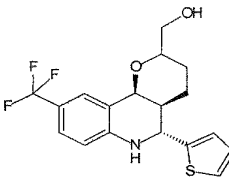


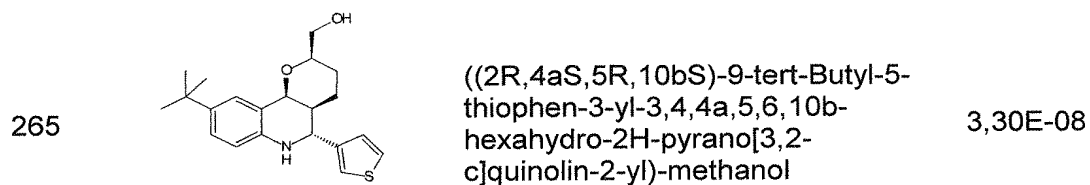
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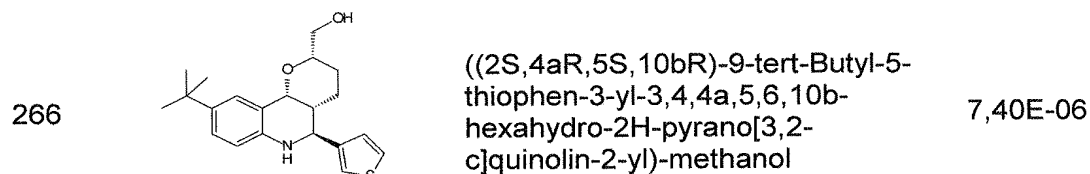
Chemistry 586

225		((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	5,67E-08
Chemistry 104			
226		((2S,4aR,5S,10bR)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	2,60E-05
Chemistry 103			
227		4-Bromo-N-(9-tert-butyl-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-N-(2-dimethylamino-ethyl)-benzamide	1,20E-05
Chemistry 102			
228		2-Fluoro-4-(2-hydroxymethyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-5-yl)-phenol	6,40E-08
Chemistry 101			
237		3-((4aS,5R,10bS)-2-Hydroxymethyl-9-methylsulfanyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-5-yl)-phenol	2,60E-07
Chemistry 289			
238		3-((4aS,5S,10bS)-2-Hydroxymethyl-9-methylsulfanyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-5-yl)-phenol	9,80E-07
Chemistry 288			
243		N-(3-Amino-propyl)-N-(9-tert-butyl-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-4-methylbenzamide	2,20E-06
Chemistry 283			

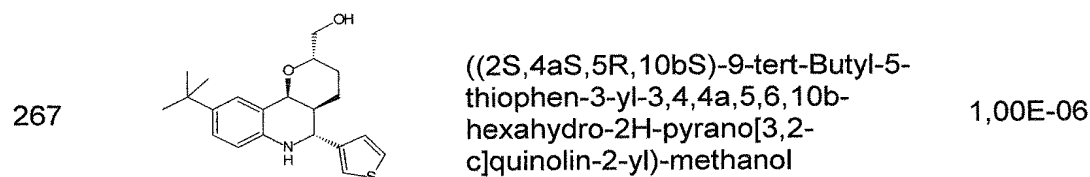
244		{3-[(9-tert-Butyl-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)methyl)-(4-methyl-benzoyl)-amino]-propyl}-carbamic acid tert-butyl ester	
	Chemistry 282		
248		((2R,4aS,5R,10bS)-9-tert-Butyl-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	8,23E-09
	Chemistry 277		
249		((2S,4aR,5S,10bR)-9-tert-Butyl-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	1,80E-05
	Chemistry 276		
250		((2R,4aR,5R,10bR)-9-tert-Butyl-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	1,70E-06
	Chemistry 275		
251		((2S,4aS,5S,10bS)-9-tert-Butyl-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	1,30E-05
	Chemistry 274		
252		((4aS,5R,10bS)-5-Thiophen-2-yl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	1,80E-07
	Chemistry 273		



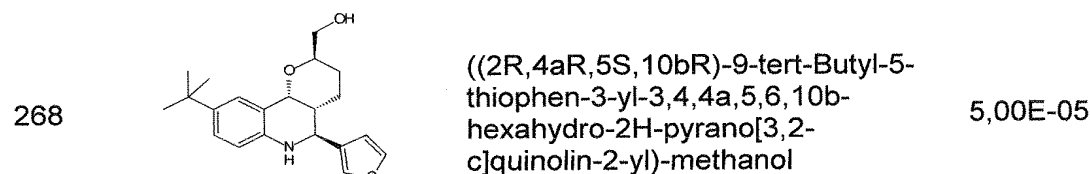
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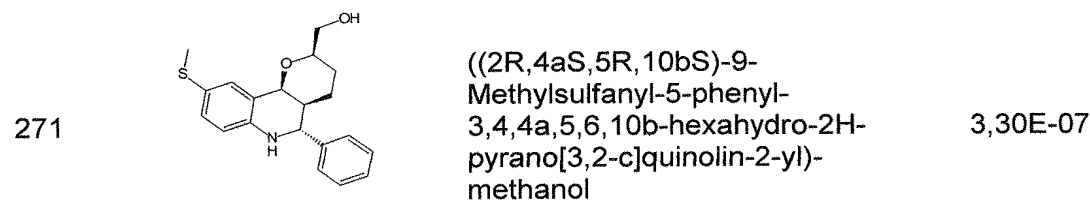
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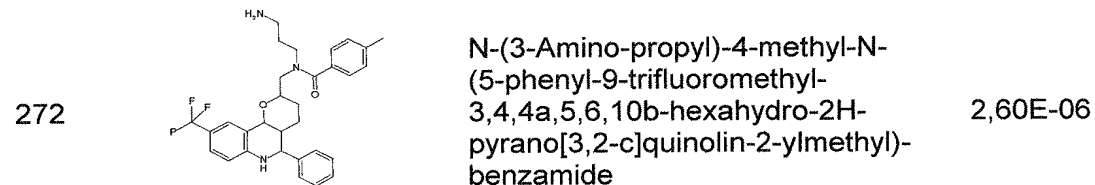
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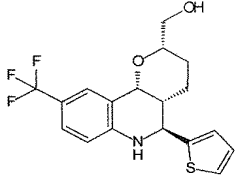
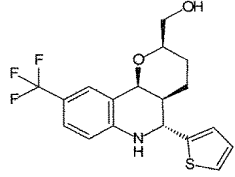
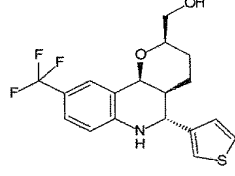
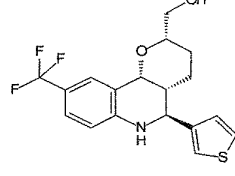
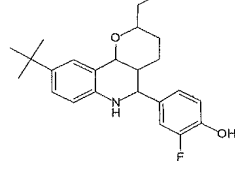
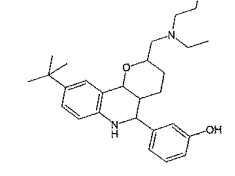
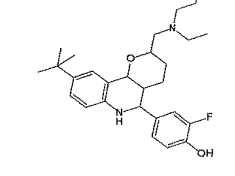
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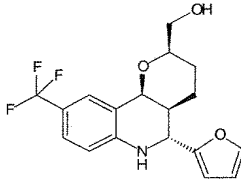
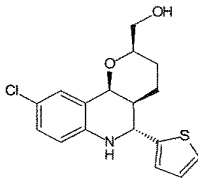
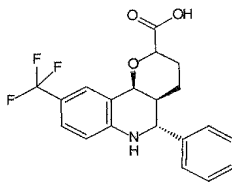
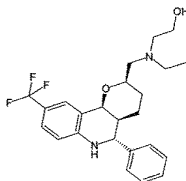
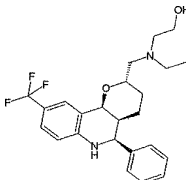
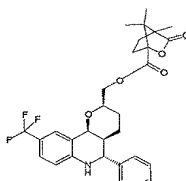


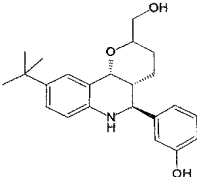
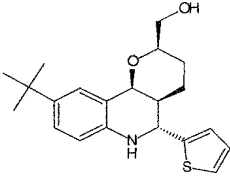
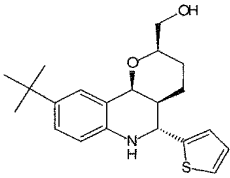
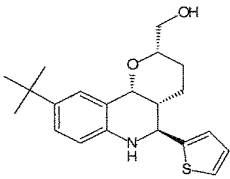
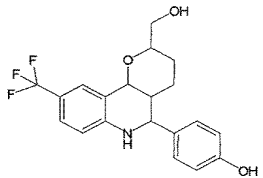
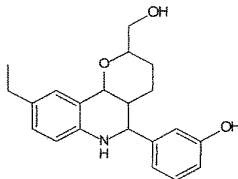
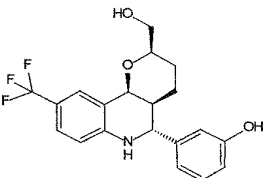
Chemistry 164

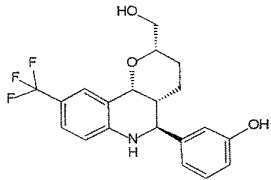
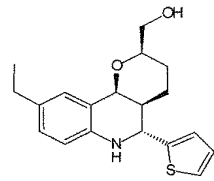
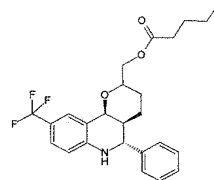
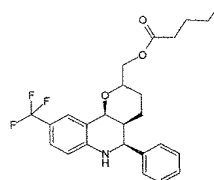
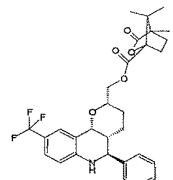
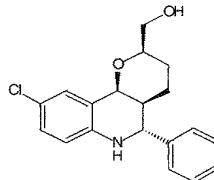


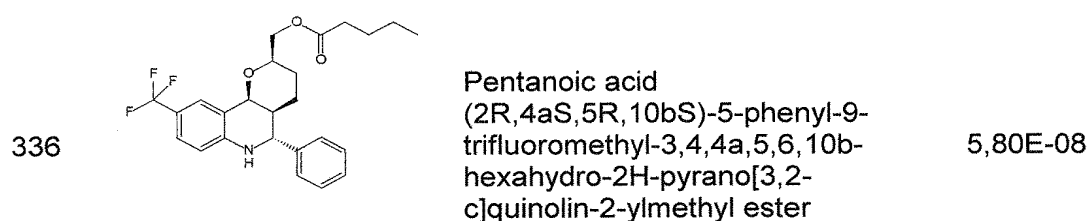
Chemistry 163

275		((2S,4aR,5S,10bR)-5-Thiophen-2-yl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	6,90E-07
	Chemistry 160		
276		((2R,4aS,5R,10bS)-5-Thiophen-2-yl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	3,10E-08
	Chemistry 159		
277		((2R,4aS,5R,10bS)-5-Thiophen-3-yl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	1,19E-07
	Chemistry 158		
278		((2S,4aR,5S,10bR)-5-Thiophen-3-yl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	
	Chemistry 157		
283		4-(9-tert-Butyl-2-hydroxymethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-5-yl)-2-fluoro-phenol	6,00E-08
	Chemistry 152		
287		3-(9-tert-Butyl-2-([ethyl-(2-hydroxy-ethyl)-amino]-methyl)-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-5-yl)-phenol	1,50E-08
	Chemistry 148		
288		4-(9-tert-Butyl-2-([ethyl-(2-hydroxy-ethyl)-amino]-methyl)-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-5-yl)-2-fluoro-phenol	2,70E-08
	Chemistry 147		

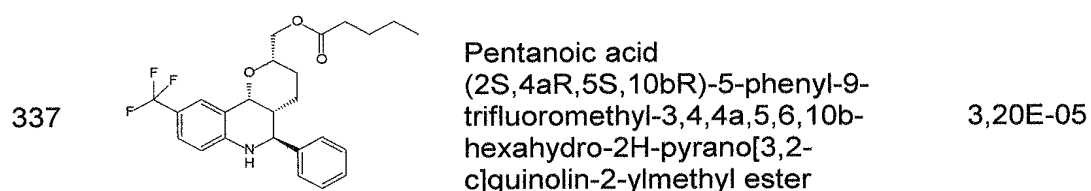
292		((2R,4aS,5R,10bS)-5-Furan-2-yl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	5,10E-07
Chemistry 143			
293		((2R,4aS,5R,10bS)-9-Chloro-5-thiophen-2-yl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	1,90E-07
Chemistry 142			
301		(4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline-2-carboxylic acid	1,10E-05
Chemistry 592			
304		2-[Ethyl-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amino]-ethanol	2,47E-07
Chemistry 593			
305		2-[Ethyl-((2S,4aS,5S,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amino]-ethanol	1,60E-05
Chemistry 594			
309		4,7,7-Trimethyl-3-oxo-2-oxa-bicyclo[2.2.1]heptane-1-carboxylic acid (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	5,10E-08
Chemistry 259			

310		3-((4aR,5S,10bR)-9-tert-Butyl-2-hydroxymethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-5-yl)-phenol	1,40E-08
	Chemistry 595		
315		((2R,4aS,5R,10bS)-9-tert-Butyl-5-thiophen-2-yl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	6,30E-09
	Chemistry 254		
319		((2R,4aS,5R,10bS)-9-tert-Butyl-5-thiophen-2-yl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	1,50E-08
	Chemistry 250		
320		((2S,4aR,5S,10bR)-9-tert-Butyl-5-thiophen-2-yl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	1,90E-06
	Chemistry 249		
323		4-(2-Hydroxymethyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-5-yl)-phenol	2,10E-07
	Chemistry 246		
324		3-(9-Ethyl-2-hydroxymethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-5-yl)-phenol	3,10E-08
	Chemistry 245		
327		3-((2R,4aS,5R,10bS)-2-Hydroxymethyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-5-yl)-phenol	4,50E-08
	Chemistry 242		

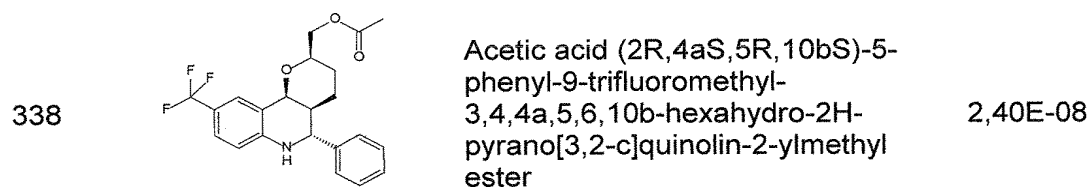
328		3-((2S,4aR,5S,10bR)-2-Hydroxymethyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-5-yl)-phenol	1,00E-05
Chemistry 241			
329		((2R,4aS,5R,10bS)-9-Ethyl-5-thiophen-2-yl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	1,80E-08
Chemistry 240			
330		Pentanoic acid (4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	3,40E-07
Chemistry 239			
331		Pentanoic acid (4aS,5S,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	1,10E-05
Chemistry 238			
334		4,7,7-Trimethyl-3-oxo-2-oxa-bicyclo[2.2.1]heptane-1-carboxylic acid (2S,4aR,5S,10bR)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	5,80E-06
Chemistry 193			
335		((2R,4aS,5R,10bS)-9-Chloro-5-phenyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methanol	9,80E-07
Chemistry 192			



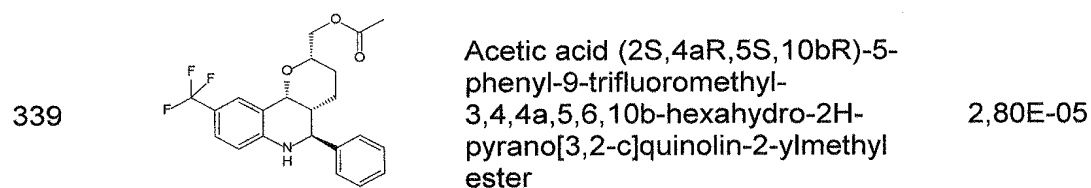
Chemistry 191



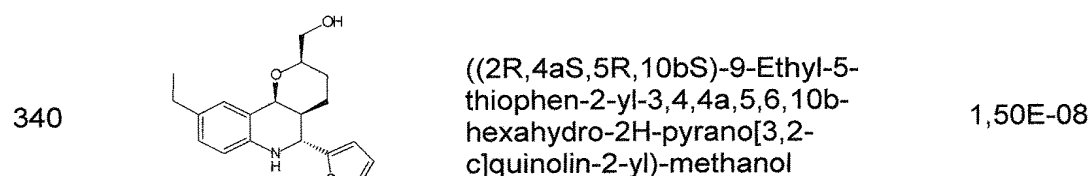
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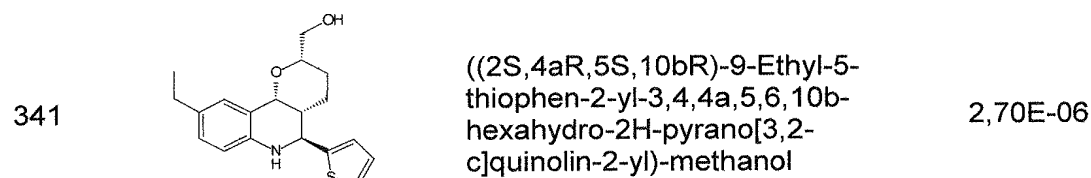
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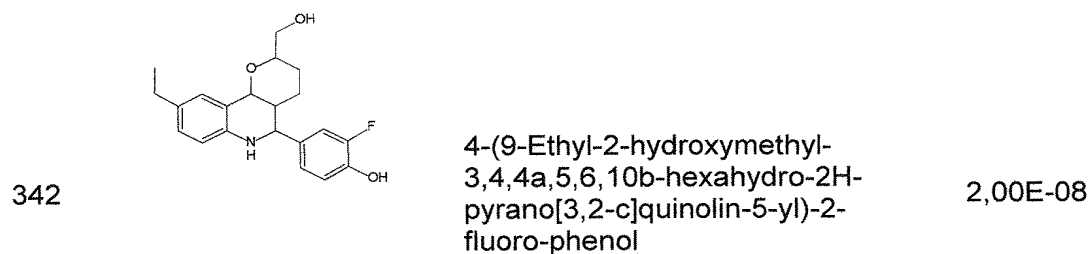
Chemistry 188



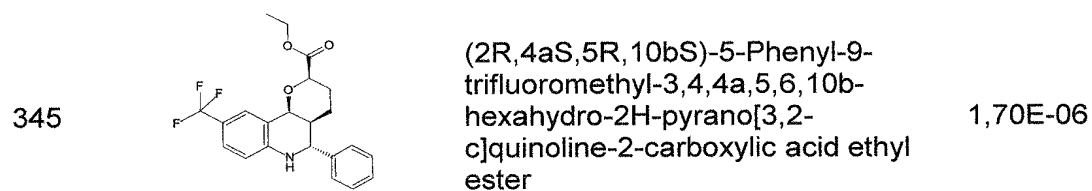
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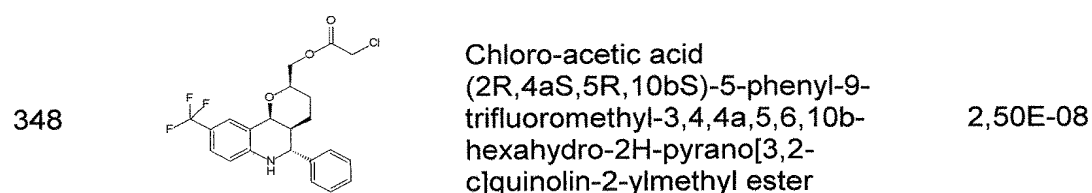
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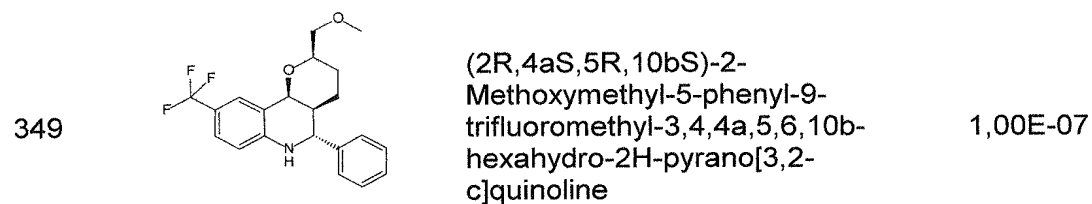
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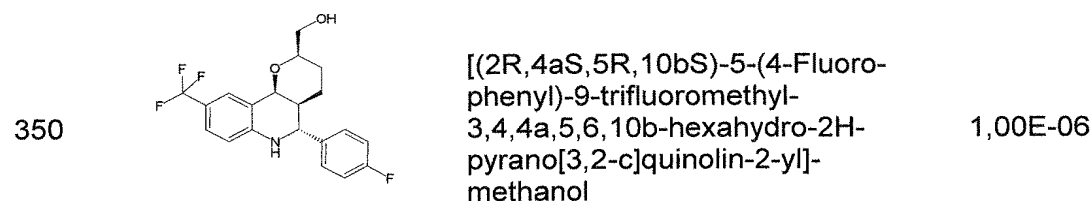
Chemistry 175



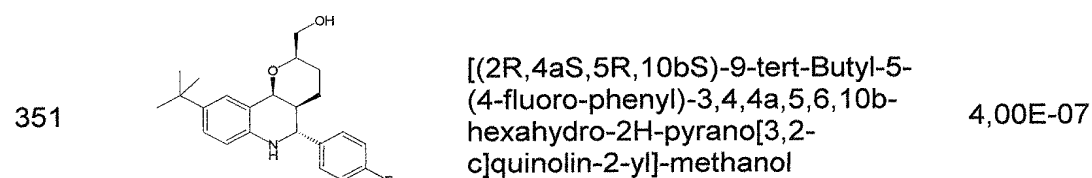
Chemistry 219



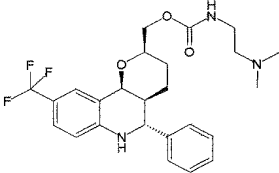
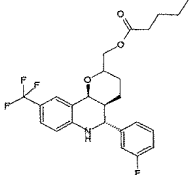
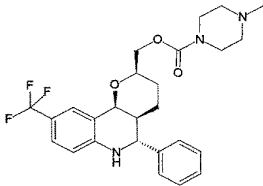
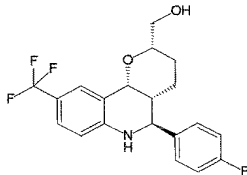
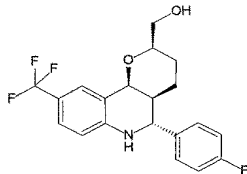
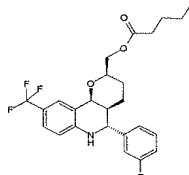
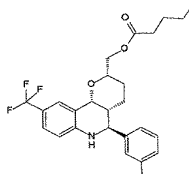
Chemistry 215



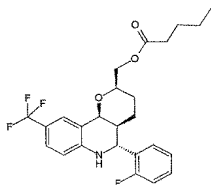
Chemistry 214



Chemistry 213

352		(2-Dimethylamino-ethyl)-carbamic acid (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	1,50E-07
	Chemistry 212		
353		Pentanoic acid (4aS,5R,10bS)-5-(3-fluoro-phenyl)-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	
	Chemistry 597		
356		4-Methyl-piperazine-1-carboxylic acid (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	2,70E-07
	Chemistry 204		
357		[(2R,4aS,5R,10bS)-5-(4-Fluoro-phenyl)-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl]-methanol	9,00E-07
	Chemistry 203		
358		[(2S,4aR,5S,10bR)-5-(4-Fluoro-phenyl)-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl]-methanol	
	Chemistry 202		
359		Pentanoic acid (2R,4aS,5R,10bS)-5-(3-fluoro-phenyl)-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	4,40E-07
	Chemistry 598		
360		Pentanoic acid (2S,4aR,5S,10bR)-5-(3-fluoro-phenyl)-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	
	Chemistry 599		

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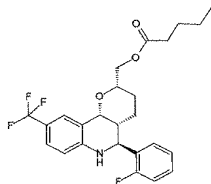


Chemistry 201

Pentanoic acid
(2R,4aS,5R,10bS)-5-(2-fluorophenyl)-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester

3,40E-07

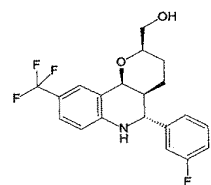
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Chemistry 200

Pentanoic acid
(2S,4aR,5S,10bR)-5-(2-fluorophenyl)-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester

363

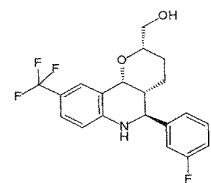


Chemistry 600

[(2R,4aS,5R,10bS)-5-(3-Fluorophenyl)-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl]-methanol

1,90E-07

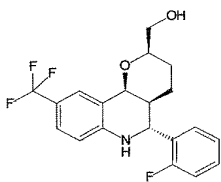
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Chemistry 601

[(2S,4aR,5S,10bR)-5-(3-Fluorophenyl)-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl]-methanol

365

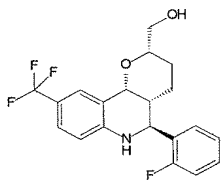


Chemistry 199

[(2R,4aS,5R,10bS)-5-(2-Fluorophenyl)-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl]-methanol

5,00E-08

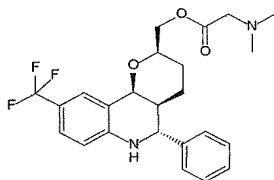
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Chemistry 198

[(2S,4aR,5S,10bR)-5-(2-Fluorophenyl)-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl]-methanol

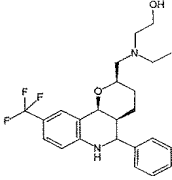
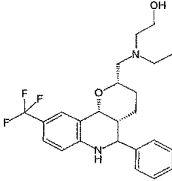
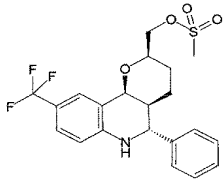
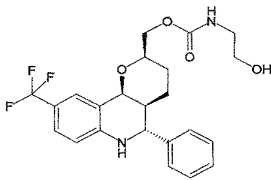
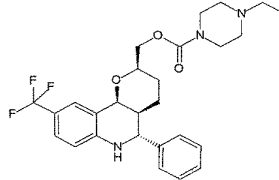
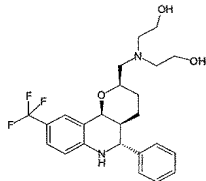
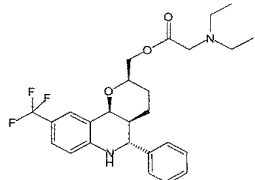
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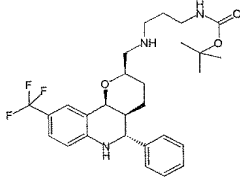
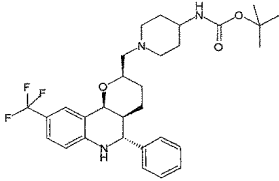
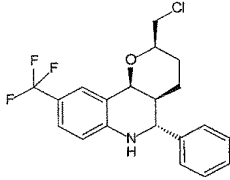
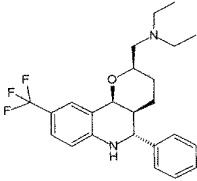
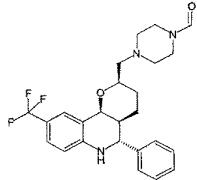
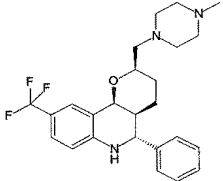
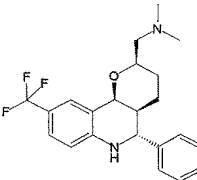


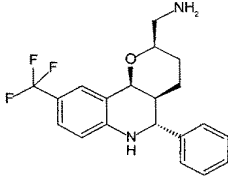
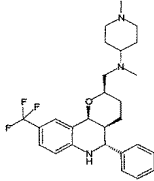
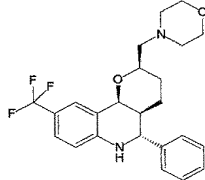
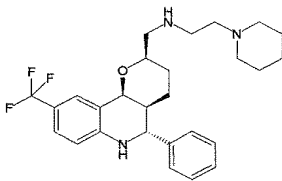
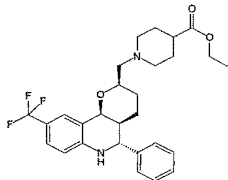
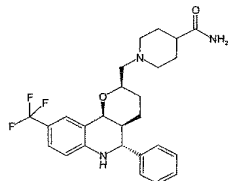
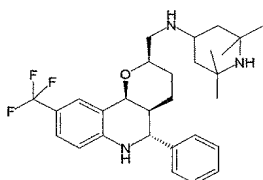
Chemistry 196

Dimethylamino-acetic acid
(2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester

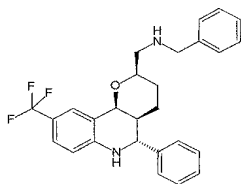
7,00E-08

562		2-[Ethyl-((2R,4aS,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amino]-ethanol	1,50E-07
563		2-[Ethyl-((2S,4aR,10bR)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amino]-ethanol	3,20E-05
564		Methanesulfonic acid (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	1,40E-07
565		(2-Hydroxy-ethyl)-carbamic acid (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	2,50E-08
566		4-Ethyl-piperazine-1-carboxylic acid (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	7,00E-08
567		2-[(2-Hydroxy-ethyl)-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amino]-ethanol	2,40E-07
568		Diethylamino-acetic acid (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	2,50E-08

569		{3-[(2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amino]-propyl}-carbamic acid tert-butyl ester	7,00E-08
570		[1-((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperidin-4-yl]-carbamic acid tert-butyl ester	2,40E-07
571		(2R,4aS,5R,10bS)-2-Chloromethyl-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	8,10E-07
572		Diethyl-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amine	3,70E-07
573		4-((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperazine-1-carbaldehyde	3,95E-07
574		(2R,4aS,5R,10bS)-2-(4-Methylpiperazin-1-ylmethyl)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	2,99E-07
575		Dimethyl-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amine	1,67E-07

576		C-((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl)-methylamine	4,01E-08
577		Methyl-(1-methyl-piperidin-4-yl)-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amine	2,93E-07
578		(2R,4aS,5R,10bS)-2-Morpholin-4-ylmethyl-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	6,04E-07
579		((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-(2-piperidin-1-yl-ethyl)-amine	1,70E-07
580		1-((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperidine-4-carboxylic acid ethyl ester	2,20E-07
581		1-((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperidine-4-carboxylic acid amide	2,50E-07
582		((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-(2,2,6,6-tetramethyl-piperidin-4-yl)-amine	2,12E-07

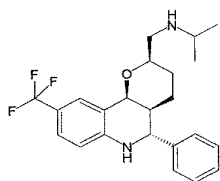
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Benzyl-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amine

3,30E-07

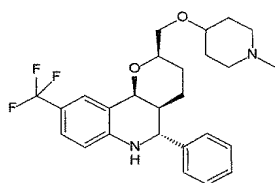
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Isopropyl-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amine

1,87E-07

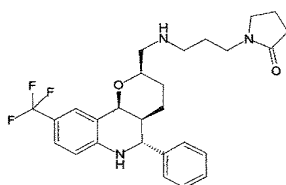
585



(2R,4aS,5R,10bS)-2-(1-Methylpiperidin-4-yloxymethyl)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline

4,10E-07

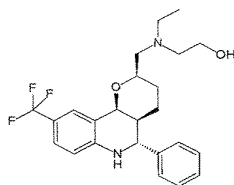
586



1-{3-[(2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amino]-propyl}-pyrrolidin-2-one

3,90E-08

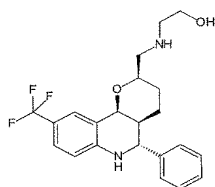
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2-[Ethyl-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amino]-ethanol

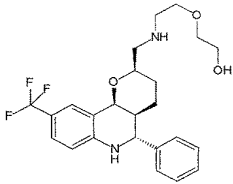
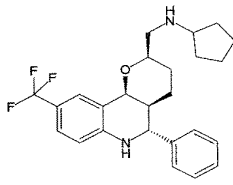
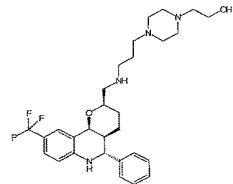
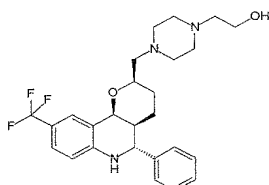
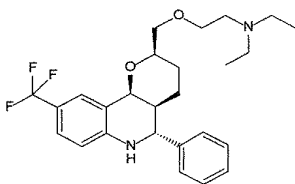
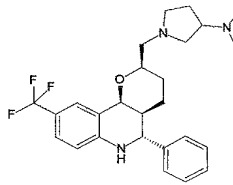
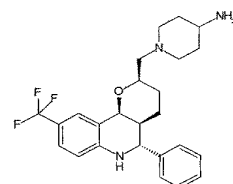
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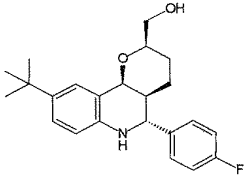
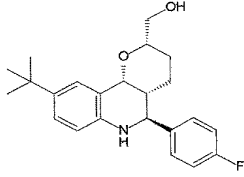
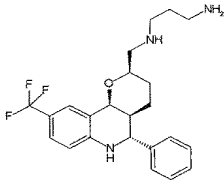
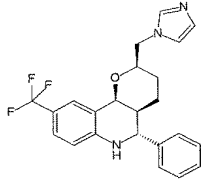
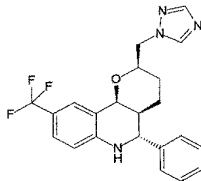
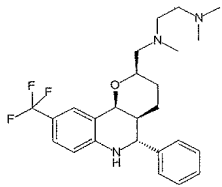
588



2-[(2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amino]-ethanol

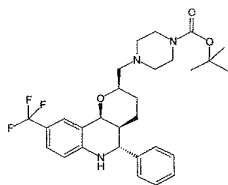
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589		2-{2-[(2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amino]-ethoxy}-ethanol	1,45E-07
590		Cyclopentyl-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amine	1,50E-07
591		2-(4-{3-[(2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amino]-propyl}-piperazin-1-yl)-ethanol	2,20E-07
592		2-[4-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperazin-1-yl]-ethanol	2,10E-07
593		Diethyl-[2-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethoxy)-ethyl]-amine	1,00E-06
594		Dimethyl-[1-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-pyrrolidin-3-yl]-amine	2,23E-07
595		1-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperidin-4-ylamine	4,40E-07

596		[(2R,4aS,5R,10bS)-9-tert-Butyl-5-(4-fluoro-phenyl)-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl]-methanol	1,90E-07
597		[(2S,4aR,5S,10bR)-9-tert-Butyl-5-(4-fluoro-phenyl)-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl]-methanol	1,00E-05
598		N1-((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-propane-1,3-diamine	8,03E-08
599		(2R,4aS,5R,10bS)-2-Imidazol-1-ylmethyl-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	2,20E-07
600		(2R,4aS,5R,10bS)-5-Phenyl-2-[1,2,4]triazol-1-ylmethyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	2,40E-08
601		N,N,N'-Trimethyl-N'-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-ethane-1,2-diamine	2,10E-07

602		(2R,4aS,5R,10bS)-5-Phenyl-2-piperazin-1-ylmethyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline	2,46E-07
603		{4-[(2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amino]-butyl}-carbamic acid tert-butyl ester	1,70E-07
604		{2-[(2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amino]-ethyl}-carbamic acid tert-butyl ester	1,60E-07
605		6-[4-((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperazin-1-yl]-pyridin-2-ol	2,00E-07
606		((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-thiophen-2-ylmethyl-amine	1,80E-07
607		4-Ethyl-1-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperidin-4-ol	7,90E-08

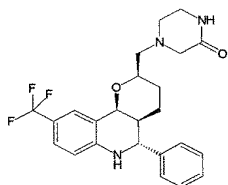
608



4-((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperazine-1-carboxylic acid tert-butyl ester

1,20E-06

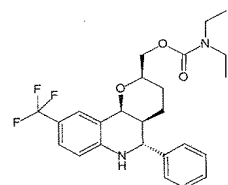
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4-((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperazine-2-one

3,20E-07

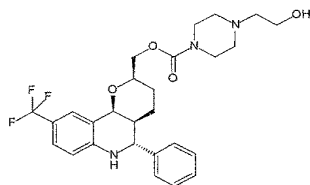
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Diethyl-carbamic acid (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester

3,40E-06

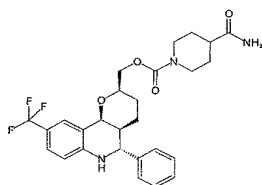
611



4-(2-Hydroxy-ethyl)-piperazine-1-carboxylic acid (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester

2,10E-07

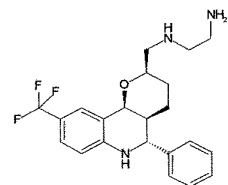
612



4-Carbamoyl-piperidine-1-carboxylic acid (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester

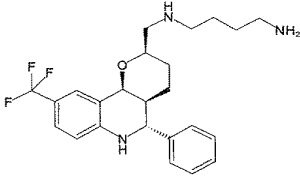
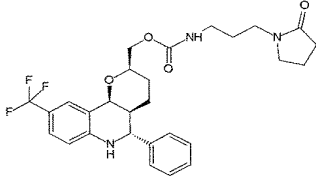
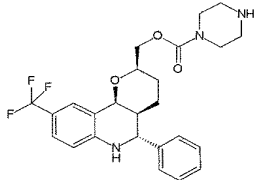
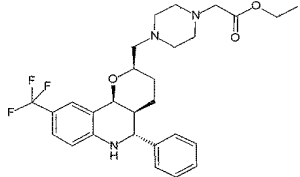
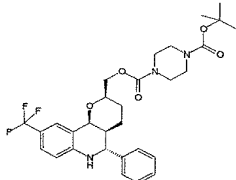
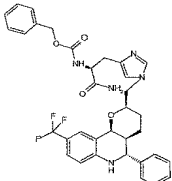
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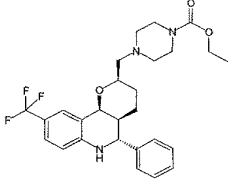
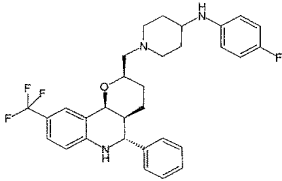
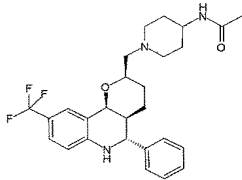
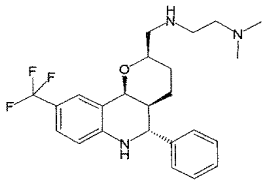
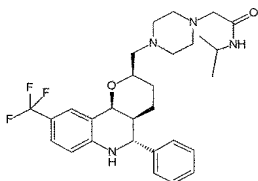
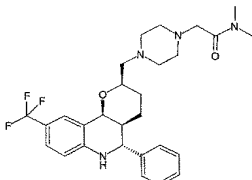
613



N1-((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-ethane-1,2-diamine

4,93E-08

614		N1-((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-butane-1,4-diamine	2,90E-07
615		[3-(2-Oxo-pyrrolidin-1-yl)-propyl]-carbamic acid (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	4,40E-07
616		Piperazine-1-carboxylic acid (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	4,00E-07
617		[4-((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperazin-1-yl]-acetic acid ethyl ester	5,30E-07
618		Piperazine-1,4-dicarboxylic acid tert-butyl ester (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	5,30E-06
619		{(S)-1-Carbamoyl-2-[1-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-1H-imidazol-4-yl]-ethyl}-carbamic acid benzyl ester	3,50E-07

620		4-((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperazine-1-carboxylic acid ethyl ester	8,80E-07
621		(4-Fluoro-phenyl)-[1-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperidin-4-yl]-amine	3,20E-07
622		N-[1-((2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperidin-4-yl]-acetamide	1,50E-07
623		N,N-Dimethyl-N'-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-ethane-1,2-diamine	4,64E-08
624		N-Isopropyl-2-[4-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperazin-1-yl]-acetamide	3,80E-07
625		N,N-Dimethyl-2-[4-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-piperazin-1-yl]-acetamide	3,30E-07

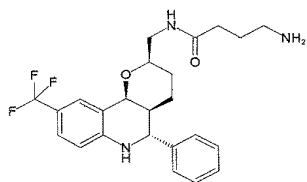
626		1-(2-Dimethylamino-ethyl)-3-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-urea	3,40E-08
627		{3-[(4-Methyl-benzoyl)-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-amino]-propyl}-carbamic acid tert-butyl ester	2,10E-06
628		Carbonic acid ethyl ester ((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	3,44E-07
629		N-(2-Amino-ethyl)-4-methyl-N-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-benzamide	2,28E-08
630		N-(4-Amino-butyl)-4-methyl-N-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-benzamide	1,70E-06
631		N-(2-Amino-ethyl)-4-methyl-N-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-benzenesulfonamide	5,00E-08

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632		N-(3-Amino-propyl)-4-methyl-N-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-benzenesulfonamide	7,50E-07
633		N-(4-Amino-butyl)-4-methyl-N-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-benzenesulfonamide	1,90E-06
634		N-(2-Amino-ethyl)-2-phenyl-N-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-acetamide	1,20E-07
635		N-(3-Amino-propyl)-2-phenyl-N-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-acetamide	3,40E-07
636		N-(4-Amino-butyl)-2-phenyl-N-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-acetamide	1,70E-06
637		1-(2-Amino-ethyl)-3-phenyl-1-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-urea	6,20E-07
638		1-(3-Amino-propyl)-3-phenyl-1-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-urea	1,60E-06

639		1-(4-Amino-butyl)-3-phenyl-1-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-urea	3,00E-06
640		N-(3-Amino-propyl)-4-methyl-N-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-benzamide	8,10E-07
641		[(2R,4aS,5R,10bS)-9-Ethyl-5-(4-fluoro-phenyl)-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl]-methanol	4,00E-07
642		(2-Amino-ethyl)-carbamic acid (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	3,90E-07
643		Carbonic acid 2-dimethylamino-ethyl ester (2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester	9,20E-08
644		2-{Ethyl-[(2R,4aS,5R,10bS)-5-(4-fluoro-phenyl)-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl]-amino}-ethanol	5,30E-07

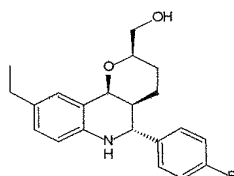
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4-Amino-N-((2R,4aS,5R,10bS)-5-phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl)-butyramide

1,70E-07

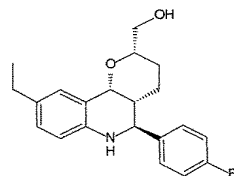
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[(2R,4aS,5R,10bS)-9-Ethyl-5-(4-fluoro-phenyl)-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl]-methanol

1,80E-07

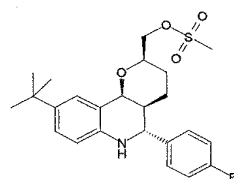
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[(2S,4aR,5S,10bR)-9-Ethyl-5-(4-fluoro-phenyl)-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-yl]-methanol

9,10E-06

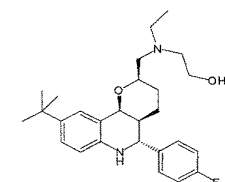
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Methanesulfonic acid (2R,4aS,5R,10bS)-9-tert-butyl-5-(4-fluoro-phenyl)-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl ester

1,10E-06

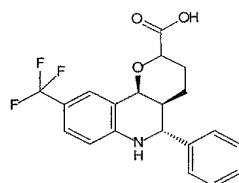
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2-([(2R,4aS,5R,10bS)-9-tert-Butyl-5-(4-fluoro-phenyl)-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinolin-2-ylmethyl]-ethyl-amino}-ethanol

1,20E-07

650

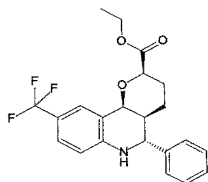


(4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline-2-carboxylic acid

1,10E-05

Chemistry 606

651

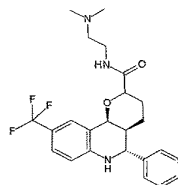


(2R,4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline-2-carboxylic acid ethyl ester

1,70E-06

Chemistry 607

652



(4aS,5R,10bS)-5-Phenyl-9-trifluoromethyl-3,4,4a,5,6,10b-hexahydro-2H-pyrano[3,2-c]quinoline-2-carboxylic acid (2-dimethylamino-ethyl)-amide

3,00E-08

Chemistry 608

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true, and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code, and that such willful false statements may jeopardize the validity of the application or any patent issuing thereon.

February 5, 2010

Date

Frank Zenke